SEQUENCE LISTING

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<110> Alitalo, Kari
      Joukov, Vladimir
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<120> VASCULAR ENDOTHELIAL GROWTH FACTOR C (VEGF-C) PROTEIN AND GENE, MUTANTS THEREOF, AND USES THEREOF

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<151> 1999-11-05

<150> PCT/US98/01973

<151> 1998-02-02

<150> 08/795,430

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<150> PCT/FI96/00427

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<170> PatentIn Ver. 2.0

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<212> DNA

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<223> Human Flt4 cDNA (short form)

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<223> Human Flt4 cDNA (3' end-long form)

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Met His Leu Leu Gly Phe Phe Ser Val Ala Cys Ser Leu Bet Ala III. 1 10 15 Ala Leu Leu Pro Gly Pro Arg Glu Ala Pro Ala Ala Ala Ala Ala Phe 20 25 30 Glu Ser Gly Leu Asp Leu Ser Asp Ala Glu Pro Asp Ala Gly Glu Ala 45
Met His Leu Leu Gly Phe Phe Ser Val Ala Cys Ser Leu Bet Alt Ala Ala Ala Ala Ala Ala Ala Leu Leu Pro Gly Pro Arg Glu Ala Pro Ala Ala Ala Ala Ala Ala Phe 20 25 30 Glu Ser Gly Leu Asp Leu Ser Asp Ala Glu Pro Asp Ala Gly Glu Ala 35 40 45 Thr Ala Tyr Ala Ser Lys Asp Leu Glu Glu Glu Gln Leu Arg Ser Val Ser
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Thr Gln Cys Met Pro Arg Glu Val Cys Ile Asp Val Gly Lys Glu Phe 130 Gly Val Ala Thr Asn Thr Phe Phe Lys Pro Pro Cys Val Ser Val Tyr 155 150 Arg Cys Gly Gly Cys Cys Asn Ser Glu Gly Leu Gln Cys Met Asn Thr Ser Thr Ser Tyr Leu Ser Lys Thr Leu Phe Glu Ile Thr Val Pro Leu 185 Ser Gln Gly Pro Lys Pro Val Thr Ile Ser Phe Ala Asn His Thr Ser 200 Cys Arg Cys Met Ser Lys Leu Asp Val Tyr Arg Gln Val His Ser Ile 215 Ile Arg Arg Ser Leu Pro Ala Thr Leu Pro Gln Cys Gln Ala Ala Asn 235 230 Lys Thr Cys Pro Thr Asn Tyr Met Trp Asn Asn His Ile Cys Arg Cys 245 Leu Ala Gln Glu Asp Phe Met Phe Ser Ser Asp Ala Gly Asp Asp Ser 265 Thr Asp Gly Phe His Asp Ile Cys Gly Pro Asn Lys Glu Leu Asp Glu Glu Thr Cys Gln Cys Val Cys Arg Ala Gly Leu Arg Pro Ala Ser Cys 290 Gly Pro His Lys Glu Leu Asp Arg Asn Ser Cys Gln Cys Val Cys Lys 315 Asn Lys Leu Phe Pro Ser Gln Cys Gly Ala Asn Arg Glu Phe Asp Glu 330 Asn Thr Cys Gln Cys Val Cys Lys Arg Thr Cys Pro Arg Asn Gln Pro 345 Leu Asn Pro Gly Lys Cys Ala Cys Glu Cys Thr Glu Ser Pro Gln Lys Cys Leu Leu Lys Gly Lys Lys Phe His His Gln Thr Cys Ser Cys Tyr 375 370 Arg Arg Pro Cys Thr Asn Arg Gln Lys Ala Cys Glu Pro Gly Phe Ser 395 390 Tyr Ser Glu Glu Val Cys Arg Cys Val Pro Ser Tyr Trp Lys Arg Pro

410

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405

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  Leu Gly Phe Ser Glu Ala Glu Pro Asp Gly Glu Val Lys Ala Phe
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                  40
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gag c Glu L	tg a eu l	atg Met 70	tct Ser	gtc Val	ctg Leu	tac Tyr	cca Pro 75	gac Asp	tac Tyr	tgg Trp	aaa Lys	atg Met 80	tac Tyr	aag Lys	tgc Cys	416
cag c Gln L	tg eu 85	cgg Arg	aaa Lys	ggc Gly	ggc Gly	tgg Trp 90	cag Gln	cag Gln	ccc Pro	acc Thr	ctc Leu 95	aat Asn	acc Thr	agg Arg	aca Thr	464
ggg g Gly <i>P</i> 100	jac Asp	agt Ser	gta Val	aaa Lys	ttt Phe 105	gct Ala	gct Ala	gca Ala	cat His	tat Tyr 110	ASII	aca Thr	gag Glu	atc Ile	ctg Leu 115	512
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Ser	gag Glu 165	gly aaa	ctg Leu	cag Gln	tgc Cys	atg Met	Asr	acc Thr	ago Sei	aca Thi	a ggt r Gly 179	A TAT	cto Lev	age 1 Se:	c aag r Lys	704 3
acg Thr 180	ttg Leu	ttt Phe	gaa Glu	att Ile	aca Thr	· Val	cct Pro	cto Lei	tca 1 Sei	a caa r Gli 19	n Gi	e eed y Pro	aaa Lys	a cc	a gto o Val 19!	•
aca Thr	atc Ile	agt Ser	ttt Phe	gcc Ala 200	a Asr	cac n His	act Thi	t too	c tg c Cy: 20	s Ar	g tg g Cy	c ato s Med	g to t Se:	t aa r Ly 21	a cto s Le 0	n 3 800
gat Asp	gtt Val	tac Tyr	aga Arg 219	g Gli	a gtt n Val	cat l His	t to	a at r Il 22	e II	t ag e Ar	a cg g Ar	t to g Se	t cter r Le 22	u FI	a gc	a 848 a
aca Thr	tta Leu	cca Pro 230	o Gli	g tg:	t cag	g gca	a gc a Al 23	a As	c aa n Ly	g ac	a tg ır Cy	t cc s Pr 24	0 111	a aa r As	ic ta sn Ty	t 896 r
gtg Val	tgg Trp 245	Ası	t aa n As:	c ta n Ty	c at r Me	g tg t Cy 25	s Ar	a tg g Cy	c ct s Le	g go u Al	et ca la Gl 25	II GI	g ga n As	t tt p Pl	t at ne Il	c 944 e
ttt Phe 260	ТУ	t tc r Se	a aa r As	t gt n Va	t ga 1 Gl 26	u As	t ga p As	c to p Se	a ac er Th	ir As	at gg sn Gl 70	ga tt Ly Ph	c ca le Hi	it ga .s As	at gt sp Va 27	• -

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aag ggg ggg ctt cgg cca tct agt tgt gga ccc cac aaa gaa cta gat 1088 Lys Gly Gly Leu Arg Pro Ser Ser Cys Gly Pro His Lys Glu Leu Asp 295 300 305
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- Glu Ser Gly Leu Gly Phe Ser Glu Ala Glu Pro Asp Gly Gly Glu Val 40
- Lys Ala Phe Glu Gly Lys Asp Leu Glu Glu Gln Leu Arg Ser Val Ser
- Ser Val Asp Glu Leu Met Ser Val Leu Tyr Pro Asp Tyr Trp Lys Met
- Tyr Lys Cys Gln Leu Arg Lys Gly Gly Trp Gln Gln Pro Thr Leu Asn
- Thr Arg Thr Gly Asp Ser Val Lys Phe Ala Ala Ala His Tyr Asn Thr
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- Leu Ser Lys Thr Leu Phe Glu Ile Thr Val Pro Leu Ser Gln Gly Pro 180
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- Ser Lys Leu Asp Val Tyr Arg Gln Val His Ser Ile Ile Arg Arg Ser 215
- Leu Pro Ala Thr Leu Pro Gln Cys Gln Ala Ala Asn Lys Thr Cys Pro 230
- Thr Asn Tyr Val Trp Asn Asn Tyr Met Cys Arg Cys Leu Ala Gln Gln 250 245
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- His Asp Val Cys Gly Pro Asn Lys Glu Leu Asp Glu Asp Thr Cys Gln 280 275

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Cys Val Cys Lys Gly Gly Leu Arg Pro Ser Ser Cys Gly Pro His Lys 290 Glu Leu Asp Arg Asp Ser Cys Gln Cys Val Cys Lys Asn Lys Leu Phe 315 Pro Asn Ser Cys Gly Ala Asn Arg Glu Phe Asp Glu Asn Thr Cys Gln Cys Val Cys Lys Arg Thr Cys Pro Arg Asn Gln Pro Leu Asn Pro Gly 345 Lys Cys Ala Cys Glu Cys Thr Glu Asn Thr Gln Lys Cys Phe Leu Lys 360 Gly Lys Lys Phe His His Gln Thr Cys Ser Cys Tyr Arg Arg Pro Cys 375 Ala Asn Arg Leu Lys His Cys Asp Pro Gly Leu Ser Phe Ser Glu Glu 395 385 Val Cys Arg Cys Val Pro Ser Tyr Trp Lys Arg Pro His Leu Asn 410 405 <210> 12 <211> 1741 <212> DNA <213> Quail <220> <221> CDS <222> (453)..(1706) <220> <223> Quail VEGF-C cDNA geceegeeg agegeteege gegeageege eggeeggge eggeeegeg agggegeget 60 <400> 12 gegageggee aetgggteet getteeetee teeteeteet teeteeteet eeteettete 120 tetgegettt ceaeegetee egagegageg eaegetegga tgteeggttt eetggtgggt 180 tttttacctg gcaaagtccg gataacttcg gtgagaattt gcaaagaggc tgggagctcc 240 cctgcaggcg tctgggagct gctgccgccg tcgcatcttc tccatcccgc ggattttact 300 gccttggata ttgcgagggg agggaggggg gtgaggacag caaaaagaaa ggggtggggg 360 gggggagaga aaaggaaaag aaggagcctc ggaattgtgc ccgcattcct gcgctgcccc 420 geggeeecce teegetetge cateteegea ca atg eac ttg etg gag atg etc

Met His Leu Leu Glu Met Leu 1

- 18 -

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gag Glu 40	gag Glu	cc Pr	c gg c G:	gt q ly <i>i</i>	gcc Ala	ggg Gly 45	gaa Glu	ccc	aa Ly	g g s A	ıct la	cat His 50	gca Ala	ago Ser	a a	aa g ys 2	gac Asp	cto Let	9 1 5	617
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cat c His H	ac ca is Gl	g aca n Thi	a tgo r Cys 380	s Sei	tgt Cys	tae Ty:	c aga	a cca g Pro 38!) PI	a tg o Cy	t ac s Th	a gt r Va	c cg l Ar 39	J	1625
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Ala His Ala Ser Lys Asp Leu Glu Glu Gln Leu Arg Ser Val Ser Ser

Val Asp Glu Leu Met Thr Val Leu Tyr Pro Glu Tyr Trp Lys Met Phe

Lys Cys Gln Leu Arg Lys Gly Gly Trp Gln His Asn Arg Glu His Ser

Ser Ser Asp Thr Arg Ser Asp Asp Ser Leu Lys Phe Ala Ala Ala His 100

Tyr Asn Ala Glu Ile Leu Lys Ser Ile Asp Thr Glu Trp Arg Lys Thr

Gln Gly Met Pro Arg Glu Val Cys Val Asp Leu Gly Lys Glu Phe Gly

Ala Thr Thr Asn Thr Phe Phe Lys Pro Pro Cys Val Ser Ile Tyr Arg 150

Cys Gly Gly Cys Cys Asn Ser Glu Gly Leu Gln Cys Met Asn Ile Ser 170

Thr Asn Tyr Ile Ser Lys Thr Leu Phe Glu Ile Thr Val Pro Leu Ser 180

His Gly Pro Lys Pro Val Thr Val Ser Phe Ala Asn His Thr Ser Cys 200

Arg Cys Met Ser Lys Leu Asp Val Tyr Arg Gln Val His Ser Ile Ile 215

Arg Arg Ser Leu Pro Ala Thr Gln Thr Gln Cys His Val Ala Asn Lys 235 225

Thr Cys Pro Lys Asn His Val Trp Asn Asn Gln Ile Cys Arg Cys Leu 250

Ala Gln His Asp Phe Gly Phe Ser Ser His Leu Gly Asp Ser Asp Thr

Ser Glu Gly Phe His Ile Cys Gly Pro Asn Lys Glu Leu Asp Glu Glu 280

Thr Cys Gln Cys Val Cys Lys Gly Gly Val Arg Pro Ile Ser Cys Gly 295 290 Pro His Lys Glu Leu Asp Arg Ala Ser Cys Gln Cys Met Cys Lys Asn 315 310 Lys Leu Leu Pro Ser Ser Cys Gly Pro Asn Lys Glu Phe Asp Glu Glu 330 Lys Cys Gln Cys Val Cys Lys Lys Thr Cys Pro Lys His His Pro Leu 345 Asn Pro Ala Lys Cys Ile Cys Glu Cys Thr Glu Ser Pro Asn Lys Cys 360 Phe Leu Lys Gly Lys Arg Phe His His Gln Thr Cys Ser Cys Tyr Arg 375 370 Pro Pro Cys Thr Val Arg Thr Lys Arg Cys Asp Ala Gly Phe Leu Leu 395 390 Ala Glu Glu Val Cys Arg Cys Val Arg Thr Ser Trp Lys Arg Pro Leu 410 Met Asn <210> 14 <211> 10 <212> PRT <213> Artificial Sequence <223> Description of Artificial Sequence: peptide <223> IgG-Kappa sequence <400> 14 Ala Val Val Met Thr Gln Thr Pro Ala Ser 5 <210> 15 <211> 22 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: oligonucleotide

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84

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<400> 47 ttttccccta gttgttacag aaga	24
<210> 48 <211> 2991 <212> DNA <213> Homo sapiens	
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sequence

<400> 48 gttttaagta gagacggggt ttcaccaacg gttgaaaata tttatcatgg tctccctaag 60 atggacggtg ttagctagga tggtctcgat ctcctgacct catgatccac ccgcctcggc 120 ctcccaaagt gctgggatta caggcgtgag ccaccgtgtc cgaccaacct taagacaaac 180 aactactgca tgattgtttt tggagacctt ttttttattc aaataaattt ttgccagcat 240 tttctgactc aaagtatagc agcaggaaga taacactttt gtgagaaaaa agtttgaata 300 cagcttactg ctgtatttaa atgaaacagt agttaatatg atattaatat attttggata 360 tattttgagt ttgttgattt tccagtcttc acccgctgct aggcctgtgg gtgttggaaa 420 tgcctgtgtt tctcaatttt gtttgcctat tagaatcctg atgtccaagc cttactccag 480 ttagaccagt taagccagaa aggcagaagg tgtactcaag catctgtttt ttcaaaatct 540 cettttgtga tgccaagtgc aatcaaagtt tagaatcatt gtaatagcaa atggttgaat 600 ggaaactcca cettetatte aaateetace ecagtetgee ettagetgtt etetttteae 660 agatctatca atgtctgaag ataactatgg caggctgatc aaatatgcat agagcaggaa 720 gacagcaaga gagtgataca ctgaccatgt tccaaatcac aaaacatctc aacaggctag 780 atcatggacc gagtetgatg ggatggaatt teataaagat acataaaaaa geatettgga 840 tacagtaaac ttaactccac aaatacaggg gaatttagac gtgactaagt agcagtacat 900 atgaaaaatt attgaggaat tttgttgact ttaagggtag tgtgagtcaa cactgtgatt 960 tggctgccag aaaataaact caatccaagg ctgtatcaac aaaggcatac tgtccattct 1020 gcatgctcat tacagcacta agtaccgagc catgttctca accgcatact tcatgaacat 1080 ggaaagctaa cagtatggtt aaggggggaa actggaactg tcatcttggg gaataaaagg 1140 gatatttagc caggagtaaa gttagcttag ggagaccatg ataaatattt tcaaaatatt 1200 tgaaggactc agttgtggaa gtgagattag atttattgtg taaaactcca ggagtcaaaa 1260 gcaatagaga gatagaagga aatgcttttc agcagtgttg ctcatcaata aagggagtga 1320 acagccacac agaatggaag gttccctgtc ctttgagata tttaagcctt caagtaaatt 1380 atgggtgagg agtttcaaat ctagagttga accagataag aaagtctctt cttccggtaa 1440 gatattatgg acctataaca tetgtgtact taaaagtaga ttgggagtga aaggcagact 1500 tttgatgttc tgtacactgt tgaaacccct tagcgtggtc ctctgtaacc tgctcaccct 1560 gececaagga ggeagetage eaatgeeace ageceaaegg aaaceceagt getttteeaa 1620 tggggaaatg cagtcacttt tetttggatg ctacacatee tttetggaat atgteteaca 1680 cacatetete titateacee cettitteaa giaaaceaae tietigeaga agetgacaat 1740 gtgtctcttt actctccacg aagattctgg cccttctctt cacctgtcag aagtttagga 1800 ttccaaaggg atcattagca tccatcccaa cagcctgcac tgcatcctga gaactgcggt 1860 tettggatea teaggeaact tteaactaca eagaceaagg gagagagggg acceeteega 1920 ggtcccatag ggttctctga catagtgatg acctttttcc aaactttgag cagggcgctg 1980 ggggccaggc gtgcgggagg gaggacaaga actcgggagt ggccgaggat aaagcggggg 2040 ctccctccac cccacggtgc ccagtttctc cccgctgcac gtggtccagg gtggtcgcat 2100 cacetetaaa geeggteeeg eeaaeegeea geeeegggae tgaaettgee eeteeggeeg 2160 cccgctcccc gcaggggaca ggggcgggga gggagagatc cagagggggg ctgggggagg 2220 tggggccgcc ggggaggagg cgagggaaac ggggagctcc agggagacgg cttccgaggg 2280 agagtgagag gggagggcag cccgggctcg gcacgctccc tccctcggcc gctttctctc 2340 acataagcgc aggcagaggg cgcgtcagtc atgccctgcc cctgcgcccg ccgccgccgc 2400 cgccgccgct cagcccggcg cgctctggag gatcctgcgc cgcggcgctc ccgggccccg 2460 cegeegeeag eegeeeegge ggeeeteete eegeeeegg caeegeegee agegeeeeeg 2520 ccgcagcgcc cgcggcccgg ctcctctcac ttcggggaag gggagggagg agggggacga 2580 cctctccaaa aagctacacc gacgcggacc geggeggegt cctccctcgc cctcgcttca 2700 cctcgcgggc tccgaatgcg gggagctcgg atgtccggtt tcctgtgagg cttttacctg 2760 acaccegeeg cettteeeeg geactggetg ggagggegee etgeaaagtt gggaacgegg 2820 agecceggae eegeteeege egeeteegge tegeceaggg ggggtegeeg ggaggageee 2880 gggggagagg gaccaggagg ggcccgcggc ctcgcagggg cgcccgcgcc cccacccctg 2940 cccccgccag cggaccggtc ccccaccccc ggtccttcca ccatgcactt g 2991

<210> 49

<211> 20

<212> DNA

<213> Artificial Sequence

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<223> Description of Artificial Sequence:
      oligonucleotide
<400> 49
                                                                    20
cacggettat gcaagcaaag
<210> 50
<211> 20
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:
      oligonucleotide
 <400> 50
                                                                     20
 aacacagttt tccataatag
 <210> 51
 <211> 19
 <212> PRT
 <213> Homo sapiens
 Leu Ser Lys Thr Val Ser Gly Ser Glu Gln Asp Leu Pro His Glu Leu
 <400> 51
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 His Val Glu
 <210> 52
 <211> 25
  <212> DNA
  <213> Artificial Sequence
  <220>
  <223> Description of Artificial Sequence:
        oligonucleotide
  <400> 52
                                                                       25
  gacggacaca gatggaggtt taaag
  <210> 53
  <211> 196
  <212> PRT
   <213> Homo sapiens
   <220>
   <223> Human PDGF-A
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His Val Leu	Ala Glu 20	Glu Ala	Glu	Ile 25	Pro 2	Arg (∃lu \	/al 1	le G 30	lu A	Arg
Leu Ala Arg 35	Ser Gln	Ile Hi	s Ser 40	Ile	Arg	Asp :	Leu (31n <i>1</i> 45	Arg L	eu 1	Leu
Glu Ile Asp 50	Ser Val	Gly Se	r Glu 5	Asp	Ser	Leu	Asp 60	Thr	Ser I	Jeu ∑	Arg
Ala His Gly 65	Val His	Ala Th	r Lys	His	Val	Pro 75	Glu	Lys	Arg I	?ro	Leu 80
Pro Ile Arg	Arg Lys	a Arg Se	r Ile	Glu	Glu 90	Ala	Val	Pro	Ala '	Val 95	Cys
Lys Thr Arg	Thr Va	l Ile Ty	r Glu	ı Ile 105	Pro	Arg	Ser	Gln	Val 110	Asp	Pro
Thr Ser Ala		e Leu I	le Trp	Pro	Pro	Cys	Val	Glu 125	Val	Lys	Arg
Cys Thr Gly	y Cys Cy	s Asn T	hr Se: 35	r Ser	· Val	Lys	Cys 140	Gln	Pro	Ser	Arg
Val His His 145	s Arg Se	r Val L 150	ys Va	l Ala	a Lys	: Val	Glu	Tyr	Val	Arg	Lys 160
Lys Pro Ly	s Leu Ly	ys Glu V 55	al Gl	n Va	1 Arg	g Lev	ı Glu	Glu	His	Leu 175	Glu
Cys Ala Cy	s Ala Ti 180	nr Thr S	er Le	u As 18	n Pro 5	o Ası	р Туг	Arg	g Glu 190	Glu	a Asp
Thr Asp Va											
<210> 54 <211> 241 <212> PRT <213> Homo	o sapier	ıs									
<220> <223> Hum	an PDGF-	-В									

 Leu Ser Asp His Ser Ile Arg Ser Phe Asp Asp Leu Gln Arg Leu Leu - 32 -His Gly Asp Pro Gly Glu Glu Asp Gly Ala Glu Leu Asp Leu Asn Met Thr Arg Ser His Ser Gly Gly Glu Leu Glu Ser Leu Ala Arg Gly Arg Arg Ser Leu Gly Ser Leu Thr Ile Ala Glu Pro Ala Met Ile Ala Glu Cys Lys Thr Arg Thr Glu Val Phe Glu Ile Ser Arg Arg Leu Ile Asp Arg Thr Asn Ala Asn phe Leu Val Trp Pro Pro Cys Val Glu Val Gln Arg Cys Ser Gly Cys Cys Asn Asn Arg Asn Val Gln Cys Arg Pro Thr Gln Val Gln Leu Arg pro Val Gln Val Arg Lys Ile Glu Ile Val Arg Lys Lys Pro Ile Phe Lys Lys Ala Thr Val Thr Leu Glu Asp His Leu Ala Cys Lys Cys Glu Thr Val Ala Ala Arg Pro Val Thr Arg Ser Pro Gly Gly Ser Gln Glu Gln Arg Ala Lys Thr Pro Gln Thr Arg Val Thr Ile Arg Thr Val Arg Val Arg Pro Pro Lys Gly Lys His Arg Lys Phe Lys His Thr His Asp Lys Thr Ala Leu Lys Glu Thr Leu Gly 225 Ala <210> 55 <211> 149 <212> PRT <213> Homo sapiens Met Pro Val Met Arg Leu Phe Pro Cys Phe Leu Gln Leu Leu Ala Gly <223> Human PIGF <220>

Leu Ala Leu Pro Ala Val Pro Pro Gln Gln Trp Ala Leu Ser Ala Gly 20

Asn Gly Ser Ser Glu Val Glu Val Val Pro Phe Gln Glu Val Trp Gly

Arg Ser Tyr Cys Arg Ala Leu Glu Arg Leu Val Asp Val Val Ser Glu

Tyr Pro Ser Glu Val Glu His Met Phe Ser Pro Ser Cys Val Ser Leu 70 65

Leu Arg Cys Thr Gly Cys Cys Gly Asp Glu Asn Leu His Cys Val Pro

Val Glu Thr Ala Asn Val Thr Met Gln Leu Leu Lys Ile Arg Ser Gly 105 100

Asp Arg Pro Ser Tyr Val Glu Leu Thr Phe Ser Gln His Val Arg Cys 120

Glu Cys Arg Pro Leu Arg Glu Lys Met Lys Pro Glu Arg Cys Gly Asp 135

Ala Val Pro Arg Arg 145

<210> 56

<211> 191

<212> PRT

<213> Homo sapiens

<220>

<223> VEGF165 precursor

Met Asn Phe Leu Leu Ser Trp Val His Trp Ser Leu Ala Leu Leu

Tyr Leu His His Ala Lys Trp Ser Gln Ala Ala Pro Met Ala Glu Gly

Gly Gln Asn His His Glu Val Val Lys Phe Met Asp Val Tyr Gln 40 35

Arg Ser Tyr Cys His Pro Ile Glu Thr Leu Val Asp Ile Phe Gln Glu 55

Tyr Pro Asp Glu Ile Glu Tyr Ile Phe Lys Pro Ser Cys Val Pro Leu

Met Arg Cys Gly Gly Cys Cys Asn Asp Glu Gly Leu Glu Cys Val Pro

Thr Glu Glu Ser Asn Ile Thr Met Gln Ile Met Arg Ile Lys Pro His 105 100

Gln Gly Gln His Ile Gly Glu Met Ser Phe Leu Gln His Asn Lys Cys 120

Glu Cys Arg Pro Lys Lys Asp Arg Ala Arg Gln Glu Asn Pro Cys Gly 135

Pro Cys Ser Glu Arg Arg Lys His Leu Phe Val Gln Asp Pro Gln Thr 155 150 145

Cys Lys Cys Ser Cys Lys Asn Thr Asp Ser Arg Cys Lys Ala Arg Gln 170

Leu Glu Leu Asn Glu Arg Thr Cys Arg Cys Asp Lys Pro Arg Arg 185

<210> 57

<211> 188

<212> PRT

<213> Homo sapiens

<220>

<223> VEGF-B167

<400> 57

Met Ser Pro Leu Leu Arg Arg Leu Leu Leu Ala Ala Leu Leu Gln Leu 10

Ala Pro Ala Gln Ala Pro Val Ser Gln Pro Asp Ala Pro Gly His Gln 25

Arg Lys Val Val Ser Trp Ile Asp Val Tyr Thr Arg Ala Thr Cys Gln 35

Pro Arg Glu Val Val Val Pro Leu Thr Val Glu Leu Met Gly Thr Val

Ala Lys Gln Leu Val Pro Ser Cys Val Thr Val Gln Arg Cys Gly Gly 70

Cys Cys Pro Asp Asp Gly Leu Glu Cys Val Pro Thr Gly Gln His Gln

Val Arg Met Gln Ile Leu Met Ile Arg Tyr Pro Ser Ser Gln Leu Gly 105

Glu Met Ser Leu Glu Glu His Ser Gln Cys Glu Cys Arg Pro Lys Lys 115

Lys Asp Ser Ala Val Lys Pro Asp Ser Pro Arg Pro Leu Cys Pro Arg 140 135 130

Cys Thr Gln His His Gln Arg Pro Asp Pro Arg Thr Cys Arg Cys Arg 150 145

Cys Arg Arg Ser Phe Leu Arg Cys Gln Gly Arg Gly Leu Glu Leu

Asn Pro Asp Thr Cys Arg Cys Arg Lys Leu Arg Arg

<210> 58

<211> 419

<212> PRT

<213> Artificial Sequence

<223> Description of Artificial Sequence: VEGF-C delta Cys156 mutant

<220>

<223> At position 156, "Xaa" can be anything other than cysteine or can be nothing

Met His Leu Leu Gly Phe Phe Ser Val Ala Cys Ser Leu Leu Ala Ala

Ala Leu Leu Pro Gly Pro Arg Glu Ala Pro Ala Ala Ala Ala Phe 25

Glu Ser Gly Leu Asp Leu Ser Asp Ala Glu Pro Asp Ala Gly Glu Ala

Thr Ala Tyr Ala Ser Lys Asp Leu Glu Glu Gln Leu Arg Ser Val Ser

Ser Val Asp Glu Leu Met Thr Val Leu Tyr Pro Glu Tyr Trp Lys Met 70

Tyr Lys Cys Gln Leu Arg Lys Gly Gly Trp Gln His Asn Arg Glu Gln

Ala Asn Leu Asn Ser Arg Thr Glu Glu Thr Ile Lys Phe Ala Ala Ala

His Tyr Asn Thr Glu Ile Leu Lys Ser Ile Asp Asn Glu Trp Arg Lys 120 115

Thr Gln Cys Met Pro Arg Glu Val Cys Ile Asp Val Gly Lys Glu Phe 135

Gly Val Ala Thr Asn Thr Phe Phe Lys Pro Pro Xaa Val Ser Val Tyr 150

Arg Cys Gly Gly Cys Cys Asn Ser Glu Gly Leu Gli Cys Not 125 165 170	Arg Cys Gly Gly Cys Cys Asn Ser Glu Gly Leu Gln Cys Me	t Asn T 175	hr
--	--	----------------	----

- Ser Thr Ser Tyr Leu Ser Lys Thr Leu Phe Glu Ile Thr Val Pro Leu 180
- Ser Gln Gly Pro Lys Pro Val Thr Ile Ser Phe Ala Asn His Thr Ser 195 200 205
- Cys Arg Cys Met Ser Lys Leu Asp Val Tyr Arg Gln Val His Ser Ile 210 215
- Ile Arg Arg Ser Leu Pro Ala Thr Leu Pro Gln Cys Gln Ala Ala Asn 225 230 235 240
- Lys Thr Cys Pro Thr Asn Tyr Met Trp Asn Asn His Ile Cys Arg Cys 245
- Leu Ala Gln Glu Asp Phe Met Phe Ser Ser Asp Ala Gly Asp Asp Ser 260 265 270
- Thr Asp Gly Phe His Asp Ile Cys Gly Pro Asn Lys Glu Leu Asp Glu 275 280 285
- Glu Thr Cys Gln Cys Val Cys Arg Ala Gly Leu Arg Pro Ala Ser Cys 290 295 300
- Gly Pro His Lys Glu Leu Asp Arg Asn Ser Cys Gln Cys Val Cys Lys 305 310 315 320
- Asn Lys Leu Phe Pro Ser Gln Cys Gly Ala Asn Arg Glu Phe Asp Glu 335
- Asn Thr Cys Gln Cys Val Cys Lys Arg Thr Cys Pro Arg Asn Gln Pro 340
- Leu Asn Pro Gly Lys Cys Ala Cys Glu Cys Thr Glu Ser Pro Gln Lys 355
- Cys Leu Leu Lys Gly Lys Lys Phe His His Gln Thr Cys Ser Cys Tyr 370 375
- Arg Arg Pro Cys Thr Asn Arg Gln Lys Ala Cys Glu Pro Gly Phe Ser 385
- Tyr Ser Glu Glu Val Cys Arg Cys Val Pro Ser Tyr Trp Lys Arg Pro 405 410 415

Gln Met Ser

<210> 59

<211> 160

<212> PRT



<213> Artificial Sequence

<223> Description of Artificial Sequence: VEGF-C delta N delta CHis

<400> 59

Met His Leu Leu Gly Phe Phe Ser Val Ala Cys Ser Leu Leu Ala Ala

Ala Leu Leu Pro Gly Pro Arg Glu Ala Pro Ala Ala Ala Ala Thr

Glu Glu Thr Ile Lys Phe Ala Ala Ala His Tyr Asn Thr Glu Ile Leu

Lys Ser Ile Asp Asn Glu Trp Arg Lys Thr Gln Cys Met Pro Arg Glu 50

Val Cys Ile Asp Val Gly Lys Glu Phe Gly Val Ala Thr Asn Thr Phe

Phe Lys Pro Pro Cys Val Ser Val Tyr Arg Cys Gly Gly Cys Cys Asn

Ser Glu Gly Leu Gln Cys Met Asn Thr Ser Thr Ser Tyr Leu Ser Lys 105 100

Thr Leu Phe Glu Ile Thr Val Pro Leu Ser Gln Gly Pro Lys Pro Val 120

Thr Ile Ser Phe Ala Asn His Thr Ser Cys Arg Cys Met Ser Lys Leu 135 130

Asp Val Tyr Arg Gln Val His Ser Ile Ile His His His His His 155 150